				<del> T</del>				·		eet 1 of 6
FORM P	TO - 14	149			ATTORNE	Y DOCK	ET NO.:	MEE-002		오
INFORM	IATION	DISCLOSURI	E STATEM	ENT	APPLICAN	TS: Mill	ler et al.			
			OIP	E ch	SERIAL NO	O.: 09/78	30,142			ECH CENTER 1600/2900
			2	3 Soas	FILING DA	\TE: Feb	oruary 9, 2	001		<u>1</u> 68
			Mr.K		EXAMINE	R. Huvn	h, P.	GROUP:	1644	<u>0/29</u>
			7074.78	AREU CENT	DOCUME					- 8
	<del>,  </del>	n o cultura Tr	DATE	NAME	DOCUME	.1413	CLASS	SUB	FILI	NG DATE IF
EXAM. INIT.		DOCUMENT NUMBER	DATE	NAME				CLASS	APPF	ROPRIATE
<u>.                                    </u>	A1	5,707.986	1/13/98	Miller et a	nl.		514	185		
-/-	A2	5,733.876	3/31/98	O'Reilly t	et al.		514	12		
	A3	5,798.349	8/25/98	Levy et al			514	185		
	NA.	5,854.205	12/29/98	O'Reilly	et al.		512	2		
	A5	Q.117.862	9/12/00	Margaron	et al.		514	185		
	A6	6,180,402	1/30/01	Granville	et al.		435	375		
	A7	6,225,303	5/1.01	Miller et	al.		514	185		
<u> </u>			FORE	IGN PATE	ENT DOCU	MENTS	<u> </u>			
EXAM. INIT.		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB CLASS	FILING DATE	ABSTR ONLY	ACT	ENGLISH LANG (Y/N)
	B1 ,	WO 95 24930 A	09'21/95							
	B2 +	WO 97 33619 A	09 18/97							
	вз •	WO 97 45450 A	12/04/97							
	B4 a	WO 98 25648 A	06 18/98							
	В5 ю	WO 99 03503 A	01/28/99							
	В6 е	WO 00 51638	09′08/00							
	В7	WO 01 51087 A	07/19/01							

EXAM. INIT.	ОТН	ER DOCUMENTS: (Including Author, Title, Date, Relevant Pages, Place of Publication)
-	Cl	Adamis et al. "Inhibition of Vascular Endothelial Growth Factor Prevents Retinal Ischemia – A

	Cl	Adamis et al. "Inhibition of Vascular Endothelial Growth Factor Prevents Retinal Ischemia – Associated Iris Neovascularization in a Nonhuman Primate" (1996) <u>Arch Ophthalmol</u> , Vol. 114, pp. 66-71.
-	C2 -	Ahmad et al "Photodynamic Therapy Results in Induction of WAF1/C1P1/P21 Leading to Cell Cycle Arrest and Apoptosis" (1998) Proc. Natl. Acad. Sci. USA, Vol. 95, pp: 6977 – 6982.
	(3	Allen et al. "Photodynamic Therapy: Tumor Targeting with Adenoviral Proteins" (1999) Photochemistry and Photobiology, Vol.70, No. 4, pp. 512-523.
	C4	Akhlynina et al. "In adin-mediated Intracellular Targeting Enhances the Photodynamic Activity of Chlorid e <sub>6</sub> " (1995) <u>Cancer Research</u> . Vol 55, pp1014-1019.

Phy N H3

12/5/02

Sheet 2 of

		Sheet 2 of 6
(5	·   :	Arenberg et al. "Interferon-y-inducible (IP-10) Is an Angiostatic Factor That Inhibits Human Normall Cell Lung Canter (NSCLC) Tumorigenesis and Spontaneous Metastases" (1996) J. Exp. Med. Vol. 184, pp 981-992.
C6	ات	Ball et al. "A Comparative Study of the Cellular Uptake and Photodynamic Efficacy of Three Novel Zinc Phthalocyanines of Differing Charge" (1999) Photochemistry and Photobiology, Vol. 69, No.3, pp: 390-39
Ox.		Berg, K. and Moan, J. "Lysosomes and Microtubules as Targets for Photochemotherapy of Cancer" (1997) Photochemistry and Photobiology, Vol. 65, No. 3, pp. 403-409.
C8	٠	Biolo et al. "Effect of Photosensitizer Delivery System and Irradiation Parameters on the Efficiency of Photodynamic Therapy of B16 Pigmented Melanoma in Mice" (1996) Photochemistry and Photobiology, Vol. 63, No. 2, pp. 224-228.
(.8	, ,	Birchler M et al. "Selective Targeting and Photocoagulation of Ocular Angiogenesis Mediated by a Phage- derived Human antibody Fragment" (1999) <u>Nature Biotechnology</u> , Vol. 17, No. 10, pp. 984-988.
CI	0	Brem et al. "The Combination of Antiangiogenic Agents to Inhibit Primary Tumor Growth and Metastasis" (1993) J. Ped. Surgery, Vol. 28, No. 10, pp. 1253-1257.
C1	11	Bressler, et al. 'Age-related Macular Degeneration' (1998) <u>Survey of Ophthalmology</u> , Vol. 32, No. 6, pp: 375-413.
CI	12	Brooks et al. "Integrin α,β <sub>3</sub> Antagonists Promote Tumor Regression by Inducing Apoptosis of Angiogenic Blood Vessels" (1994) <u>Cell.</u> Vol. 79, pp: 1157-1164.
Cl	13	Brooks et al. "Requirement of Vascular Integrin α <sub>3</sub> β <sub>3</sub> for Angiogenesis" (1994) <u>Science</u> , Vol. 264, pp. 569-571.
C	14	Ciardiello et al. "Antitumor Activity of Combined Blockade of Epidermal Growth Factor Receptor and Protein Kinase A" (1996) <u>J. Natl. Can. Inst.</u> Vol. 88, No. 23, pp: 1770-76.
	15	Ciulla T A et al. "Changing Therapeutic Paradigms for Exudative Age-related Macular Degeneration: Antiangiogenic Agents and Photodynamic Therapy" (1999) Expert Opinion on Investigational Drugs, Ashley Publications LTD, Vol. 8, No. 12, pp. 2173-2182.
С	16	D'Amato et al "Thalidomide is an Inhibitor of Angiogenesis" (1994) Proc. Natl. Acad. Sci. USA, Vol. 91, pp. 4082-4085.
C	17	Dimitroff C J et al. "Anti-Angiogenic Activity Of Selected Receptor Tyrosine Kinase Inhibitors, PD166285 And PD173074: Implications For Combination Treatment With Photodynamic Therapy" (1999) Investigational New Drugs, Vol. 17, No. 2, pp. 121-35.
	18	Dobi et al. "A New Model of Experimental Choroidal Neovascularization in the Rat" (1989) Arch Ophthalmol, Vol. 107, pp: 264-269.
C	19,	Dougherty et al. "Photodynamic Therapy" (1998) J. Natl. Can. Inst., Vol. 90, No. 12, pp: 889-905.
C	20	Ellerby et al. "Anti-cancer activity of targeted pro-apoptotic peptides" (1999) Nature Medicine, Vol. 5, No. 9, pp: 1032-1038.
C	21	Ferrario et al. "Antiangiogenic Treatment Enhances Photodynamic Therapy Responsiveness in a Mouse Mammary Carcinoma" (2000) Cancer Research, Vol. 60, No. 15, pp. 4066-4069.
(	`22 •	Fingar et al. "Analysis of Acute Vascular Damage After Photodynamic Therapy Using Benzoporphyrin Derivative (BPD)" (1999) <u>British Journal of Cancer</u> , Vol. 79, pp. 1702-1708.
(	'23	Folkman, J. "Angiogenesis in Cancer, Vascular, Rheumatoid and Other Disease" (1995) Nature Medicine, Vol. 1, pp. 27-31.
	24	Friedlander et al. "Definition of Two Angiogenic Pathways by Distinct α <sub>v</sub> Integrins" (1995) <u>Science</u> . Vol. 270, pp. 1500-1502.
L		

Py of

13/5/02

APR	2	9 2002	F
A.C.		, ask	2000

C25	Friedlander et al. "Involvement of Integrins $\alpha_{\nu}\beta_3$ and $\alpha_{\nu}\beta_5$ in Ocular Neovascular Diseases" (1996) Pro. N Acad. Sci. USA, Vol. 93, pp. 9764-9769.
C26	Granville et al. "Overexpression of Bcl-X <sub>L</sub> Prevents Caspase-3-mediated Activation of DNA Fragmentation (DFF) Produced by Treatment with the Photochemotherapeutic Agent BPD-MA" (1998) <u>FEBS</u> <u>Letters</u> , Vol. 422, pp. 151-154.
(2)	Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31 Cleavage in HeLa cells Treated with Photodynamic Therapy" (1998) FFBS Letters, Vol. 437, pp. 5-10.
C28	Oupta et al. "Involvement of Nitric Oxide During Phthalocyanine (Pc4) Photodynamic Therepy-Mediated Aportosis" (1998) Cancer Research, Vol. 58, pp. 1785-1788.
C29	Guyer et al. "Subfoveal Choroidal Neovascular Membranes in Age-Related Macular Degeneration" (198 Arch Ophhalmol. Vol. 104 pp: 702-705.
C30	Hakem et al. Differential Requirement for Caspase 9 in Apoptotic Pathways in Vivo" (1998) Cell, Vol. pp: 339-352.
C31	Hanahan, D. and Folkman, J. "Patterns and Emerging Mechanisms of the Angiogenic Switch during Tumorigenesis" (1996 Cell, Vol. 86, pp. 353-364.
C32	He et al. "Photodynamic Therapy with Photofrin II Induces Programmed Cell Death in Carcinoma Cell Lines" (1994) Photochemistry and Photobiology, Vol. 59, No. 4, pp. 468-473.
C33	He et al. "The Induction of Partial Resistance to Photodynamic Therapy by the Protooncogene <i>BCL-2</i> " (1996) <u>Photochemistry and Photobiology</u> . Vol. 64, No. 5, pp. 845-852.
C34	He et al. Variation in Photodynamic Efficacy during the Cellular Uptake of Two Phthalocyanine Photosensitizers" (1998) Photochemistry and Photobiology, Vol. 67, No. 6, pp. 720-728.
C35	A Provident Pathyras to Provent Apontosis" (1993) Cell. Vol.
C36	Hunt David W.C. et al. "Impact of PDT and Fas-mediated Apoposis on Cells of the System" (1998)  Photochemistry and Photobiology, Vol. 67, Spec. Issue, pp. 698.
C37	Husain et al. "Photodynamic Therapy." (Age-Related Macular Degeneration). Chapter 17, pp. 297-307
ļ.	Husain, et al. 'Intravenous Infusion of Liposomal Benzoporphyrin Derivative for Photodynamic Therap Experimental Choroidal Neovascularization" (1996) <u>Arch Ophthalmol</u> , Vol. 114, pp. 978-985.
C39	Epidemiology, Vol. 118, No. 2., pp: 213-227.
C40	V 1 2 19 mm 555 557
C4	Kane et al. "Bel-2 Inhibition of Neural Death: Decreased Generation of Reactive Oxygen Species" (19 Science, Vol. 262, No. 19, pp. 1274-1277.
C4	Kerr, et al. "Apoptosis: A Basic Biological Phenomenon with Wide-Ranging Implications in Tissue Kinetics" (1972) Br. J. Cancer, Vol. 26, pp. 239-257.
C4	Kessel et al. Biodistribution of Photosensitizing Agents" (1993) Int. J. Biochem, Vol. 25, No. 10, pp: 83.
C4	<ul> <li>Kessel, D. and Luo, Y "Photodynamic therapy: A Mitochondrial Inducer of Apoptosis" (1999) Cell Inducer of Apoptosis (19</li></ul>
C4	<ul> <li>Klein, B. and Klein, R."Cataracts and Macular Degeneration in Older Americans" (1982) <u>Arch Ophthe</u></li> <li>Vol. 100, pp; 571-573.</li> </ul>

APR 2 9 2002 8

	80,142	Sheet 4
	C46	Kramer, et al. "Liposomal Benzoporphyrin Derivative Verteporfin Photodynamic Therapy" (1996)  Ophthalmology Vol. 103, No. 3, pp. 427-438.
/	C47	Kramer, et al. "Liposomal Benzoporphyrin Derivative Verteporfin Photodynamic Therapy" (1996)  Ophthalmology Vol. 103, No. 3, pp: 427-438.  Kroemer, G. "The Proto-oncogene Bcl-2 and Its Rolc in Regulating Apoptosis" (1997) Nature Medicine, Vol. 3, No. 6, pp: 614-620.  Lucas et al. "Multiple Forms of Angiostatin Induce Apotosis in Endothelial Cells" (1998) Blood, Vol. 92, No. 12, pp: 4730-4741.
	C48	Lucas et al. "Multiple Forms of Angiostatin Induce Apotosis in Endothelial Cells" (1998) <u>Blood</u> , Vol. 92, No. 12, pp. 4730-4741.
	C49	Luo et al. "The Role of Mitochondrial Photodamage in PDT-induced Apoptosis" (1998) Chemical Abstract Vol. 129, No. 13 & Proc. Spie-Int. Soc. Opit. Eng. Vol. 3247, pp: 112-117.
	C50	Manceri et al. "Combined effects of Angiostatin and Ionizing Radiation in Antitumour Therapy" (1998)  Nature Vol. 394, pp. 287-291.
	C51	Miller et at "Alternative Treatments of Neovascular Age-Related Macular Degeneration" (1999) Chapter 139, pp 2014-2027.
	C52	Miller JW et al. Photodynamic Therapy of Experimental Chorodial Neovascularization Using Lipoprotei deliver Benzoporphyrin' (1995) <u>Archives of Opthalmology</u> , Vol. 113, No. 6, pp. 810-818.
	C53	Mori et al. "Photodynamic Therapy for Experimental Tumors Using ATX-S10(Na), a Hydrophilic Chlorin Photosensitizer, and Diode Laser." (2000) <u>Jpn. J. Cancer</u> , Vol. 91, No. 7, pp: 753-9.
	C54	Moser et al. "Angiostatin binds ATP Synthase on the Surface of Human Endothelial Cells" (1999) Proc. N Acad. Sci USA, Vol. 96, pp. 2811-2816.
	C55	Mosmann, T "Rapid Colorimetric Assay for Cellular Growth and Survival: Application to Proliferation ar Cytotoxicity Assays" (1983) Journal of Immunological Methods, Vol. 65, pp. 55-63.
	C56	Netland et al. In situ Labelling of Vascular Endothelium with Fluorescent Acetylated Low Density Lipoprotein" (1985) <u>Histochem L.</u> Vol. 17, pp. 1309-1320.
	C57	Nicholson, D. and Thornberry, N. "Capases: Killer Proteases" (1997) TIBS, Vol. 22, pp. 299-306.
	C58	Obana et al. "Selective Occlusion of Choroidal Neovascularization by Photodynamic Therapy with a Water Soluble Photosensitizer, ATX-S10" (1999) <u>Lasers Sura, Med.</u> , Vol. 24, No. 3, pp. 209-222.
	C59	Obana et al. "Selective Photodynamic Effects of the New Photosensitizer ATX-S10(Na) on Choroidal Neovascularization in Monkeys" (2000) <u>Arch. Ophthalmol.</u> Vol. 118, No. 5, pp. 650-658.
	C60	Okamoto et al. "Animal Model: Transgenic Mice with Increased Expression of Vascular Endothelial Grov Factor in the Retina" (1997) <u>Amer. J. Path.</u> Vol. 151. No. 1, pp: 281-291.
	C61	Olcinick et al. "The Photobiology of Photodynamic Therapy: Cellular Pargets and Mechanisms" (1998), Radiation Research, 150 (Suppl.), pp. S146-S156.
	C62	(00.402
	C63	O'Reilly et al. "Angiostatin: A Novel Angiogenesis Inhibitor That Mediates the Superession of Metastasco by a Lewis Lung Carcinoma" (1994) Cell. pp: 315-328.
	C64	O'Reilly et al. "Endostatin: An Endogenous Inhibitor of Angiogenesis and Tumor Growth" (1997) Cell. V 88, pp. 277-285.
	C65	Pidgeon & Colles et al. "Recent Developments In Tunable Lasers For Spectroscopy" (1979) Nature Vol. 279, pp: 377-381.
<del> </del>	C66	Pollack et al. "Repair of Retinal Pigment Epithelium and Choriocapillaries After Laser Photocoagulation Correlations Between Scanning Electron, Transmission Electron and Light Microscopy" (1997) Opthal. J Vol. 29, pp. 393-404.

C67	Reddi, E. "Role of Delivery Vehicles for Package Mozers in the Photodynamic Therapy of Tumors" (1997)  J. Photochem & Photobiol. Vol 37, pp 189-195.
C68	Renno et al. "Photodynamic Therapy Using Lu-Tex Induces Apoptosis In Vitro, and its Effect is Potentiated by Angiostatin in Retinal Capillary Endothelial Cells" (2000) Inves. Ophthal. & Vis. Sci., Vol. 41. No. 12, pp. 3963-71.
 120	Ryan "Subretinal Neovascularization: Natural History of an Experimental Model" (1982) <u>Arch Ophthalmol.</u> Vol. 100, pp: 1804-1809.
(270	Schmidt-Erfurth, U. et al. "Photodynamic therapy in Ocular Vascular Disease" (1998) <u>Laser Phys.</u> , Vol. 8, No. 1, pp; 191-198.
 C71	Senior K. "Ageing eyes retain their mystery." (1999) The Lancet Vol. 353, pp. 818.
C72	Separovic et al "Association of Ceramide Accumulation with Photodynamic Treatment-Induced Cell Death" (1998) Photochemistry and Photobiology, Vol. 68, No. 1, pp. 101-109.
C73	Sharma A. et al. "Combination of Photodynamic Therapy and Angiogenic Therapy for the Treatment of Human Breast Cancer in Nude Athymic Myce" (1996) Proceedings of the Annual Meeting of the American Association for Cancer Research, US, Phildelphia, AACR, Vol. Meeting 87, pp. 291.
C74	Stellmach et al. "Prevention of Ischemia-Induced Retinopathy by the Natural Ocular Antiangiogene Agent Pigment Epithelium-Derived Factor" (2000) Proc. Natl. Acad. Sci. pp: 1-5.
C75	Tolentino et al. "Vascular Endothelial Growth Factor Is Sufficient to Produce Iris Neovascularization and Neovascular Glaucoma in a Nonhuman Primate" (1996) <u>Arch Ophthalmol.</u> Vol. 114, pp. 964-969.
C76	Varnes et al. "Photodynamic Therapy Induced Apoptosis in Lymphoma Cells: Translocation of Cytochrome e Causes Inhibition of Respiration as Wall as Caspase Activation" (1999) <u>Biochem. &amp; Biophys. Res. Com.</u> , Vol. 255, pp. 673-679.
C77	Veis, et al. "Bc1-2-Deficient Mice Demonstrate Fulminant Lymphoid Apoptosis, Polycystic Kidneys, and Hypopigmented Hair" (1993) Cell, Vol. 75, pp. 229-240.
C78	16 midules To Ovidative Stress (1998) Free Radical Biology & Medicine, Vol. 24,
C79	Voest et al. "Inhibition of Angiogenesis In Vivo by Interleukha 12" (1995) J. Nat. Can. Inst. Vol. 87, No. 8, pp. 581-586.
C80	Antibodies: A New Possibility for Use in Photodynamic Therapy" (2000) Int. J. Cancer, Vol. 88, pp. 108-114.
C81	The World Con Page Vol 36 pp. 2326-2329
C82	C. H. L. Difference Profife ration States" (1990) Int. I. Radiat. Biol., Vol. 58, No. 1, pp. 145-156.
C83	0123) in the Murine EMT6 Sarcoma Model" (1997) Photochemistry and Photobiology, Vol. 65, No. 3, pp: 410-415.
(*8-	Photochemistry and Photobiology, Vol. 63, No. 6, pp. 892-897.
C8:	Zhang et al. "Antisense Bc1-2 Retrovirus Vector Increases the Sensitivity of a Human Gastric Adenocarcinoma Cell Line to Photodynamic Therapy" (1999) <u>Photochemistry and Photobiology</u> , Vol. 69. No. 5, pp. 582-586.

Phy & He

12/5/02

Sheet 6 of 6

DATE CONSIDERED (A)
<b>}</b>